

B

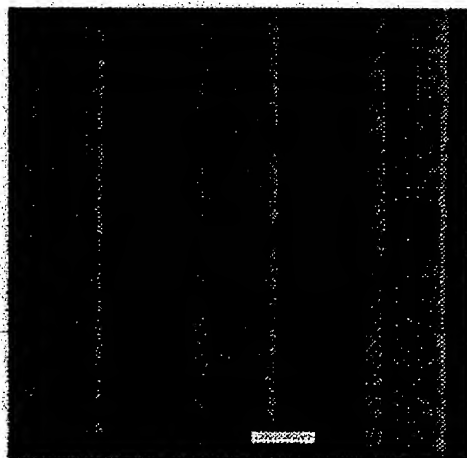


FIG. 1

2/13

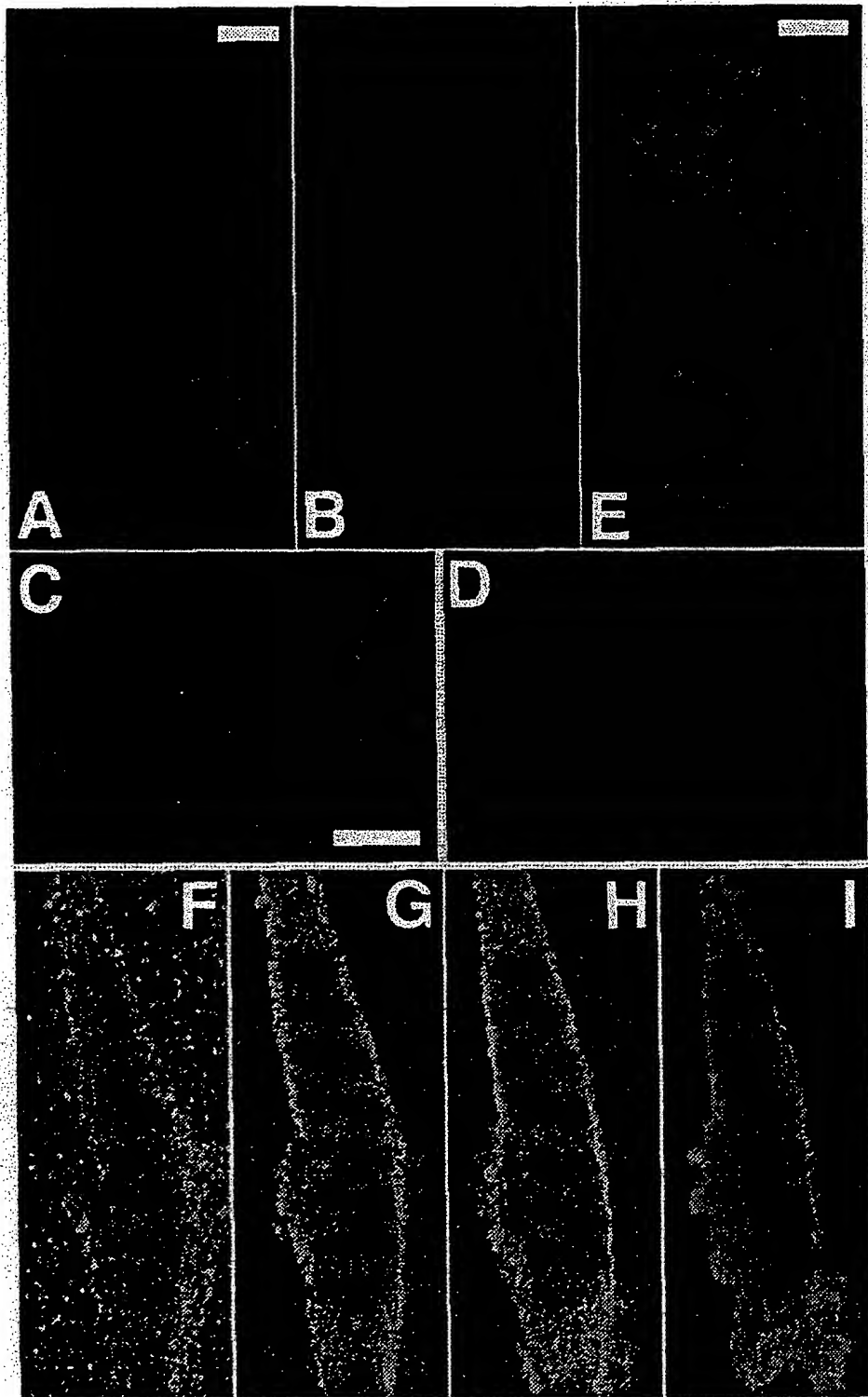


FIG. 2

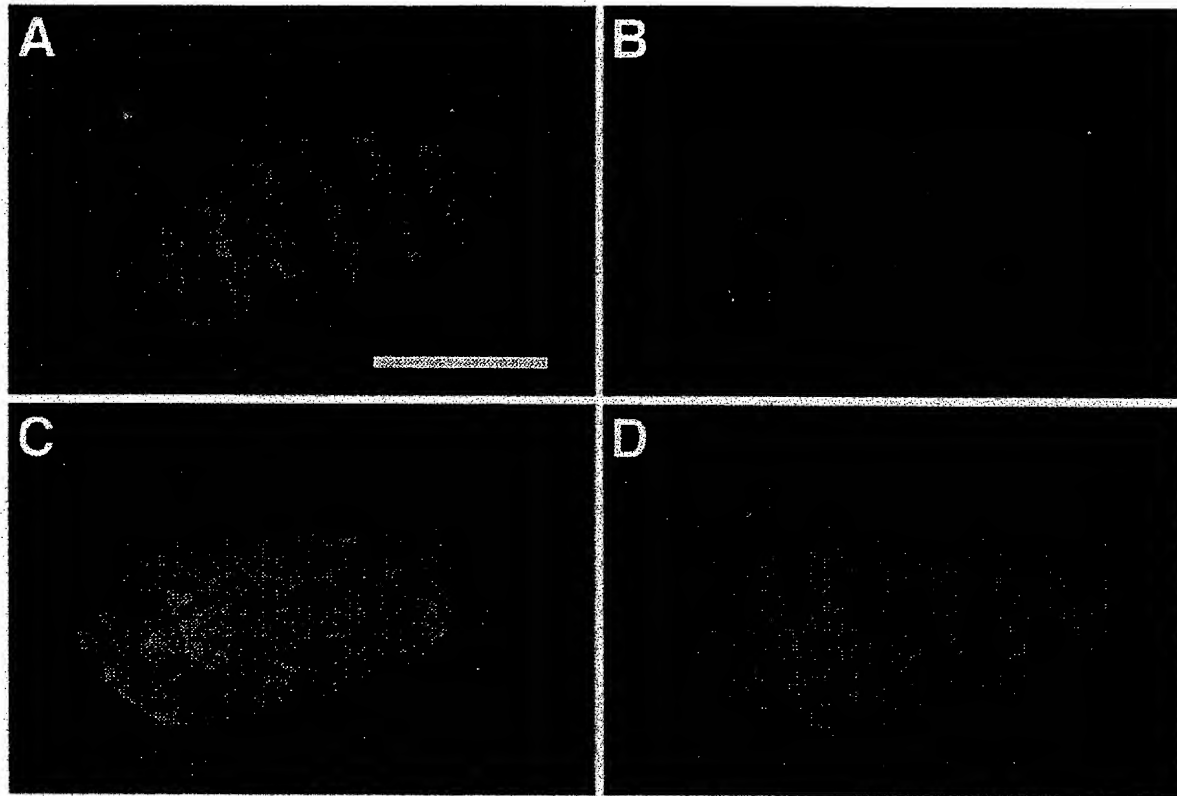


FIG. 3

4/13

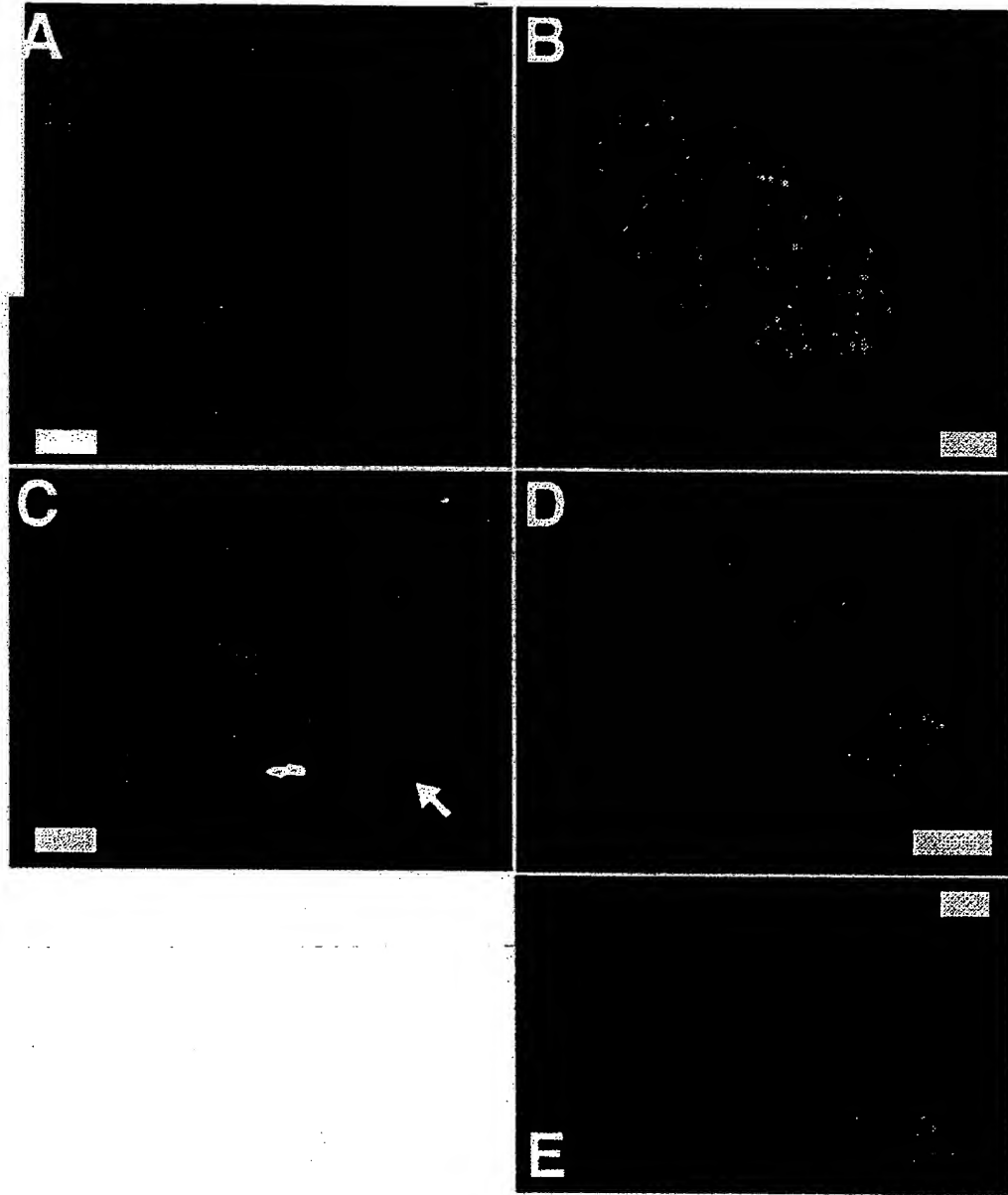


FIG. 4

5/13

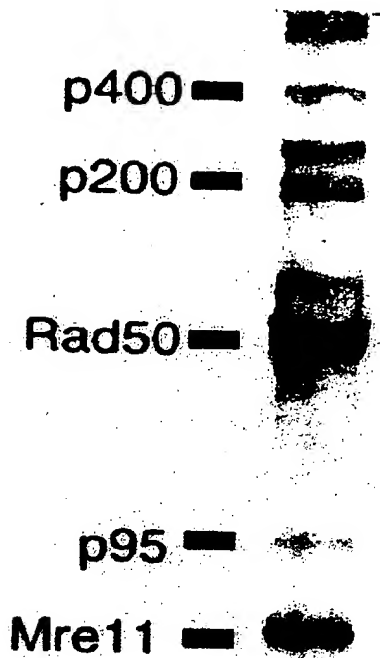


FIG. 5

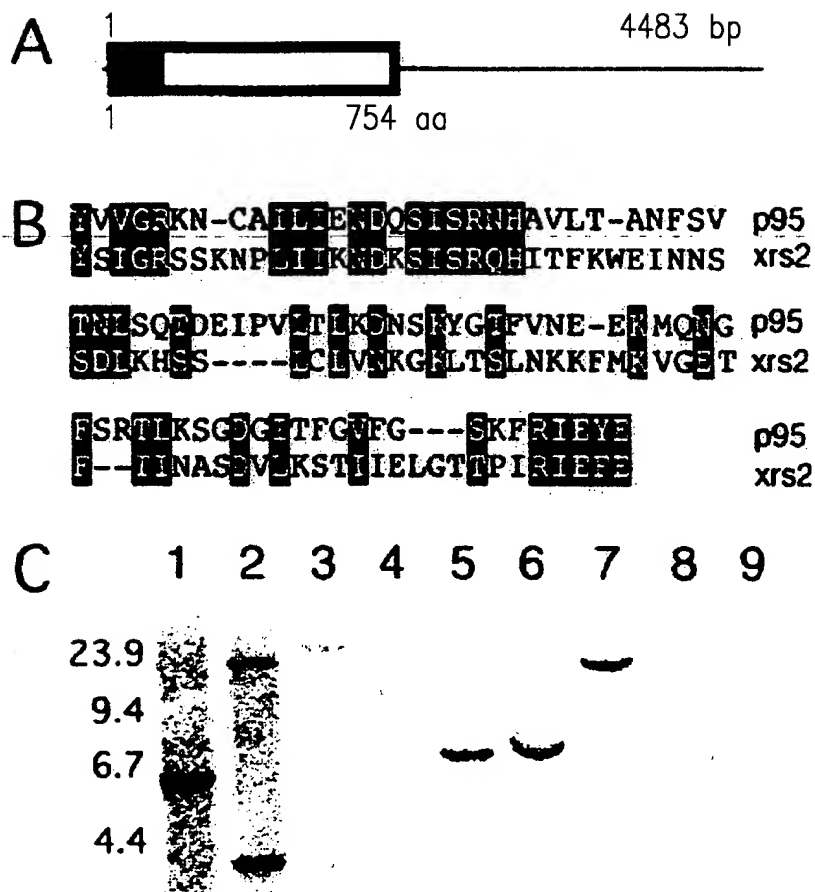


FIG. 6

6/13

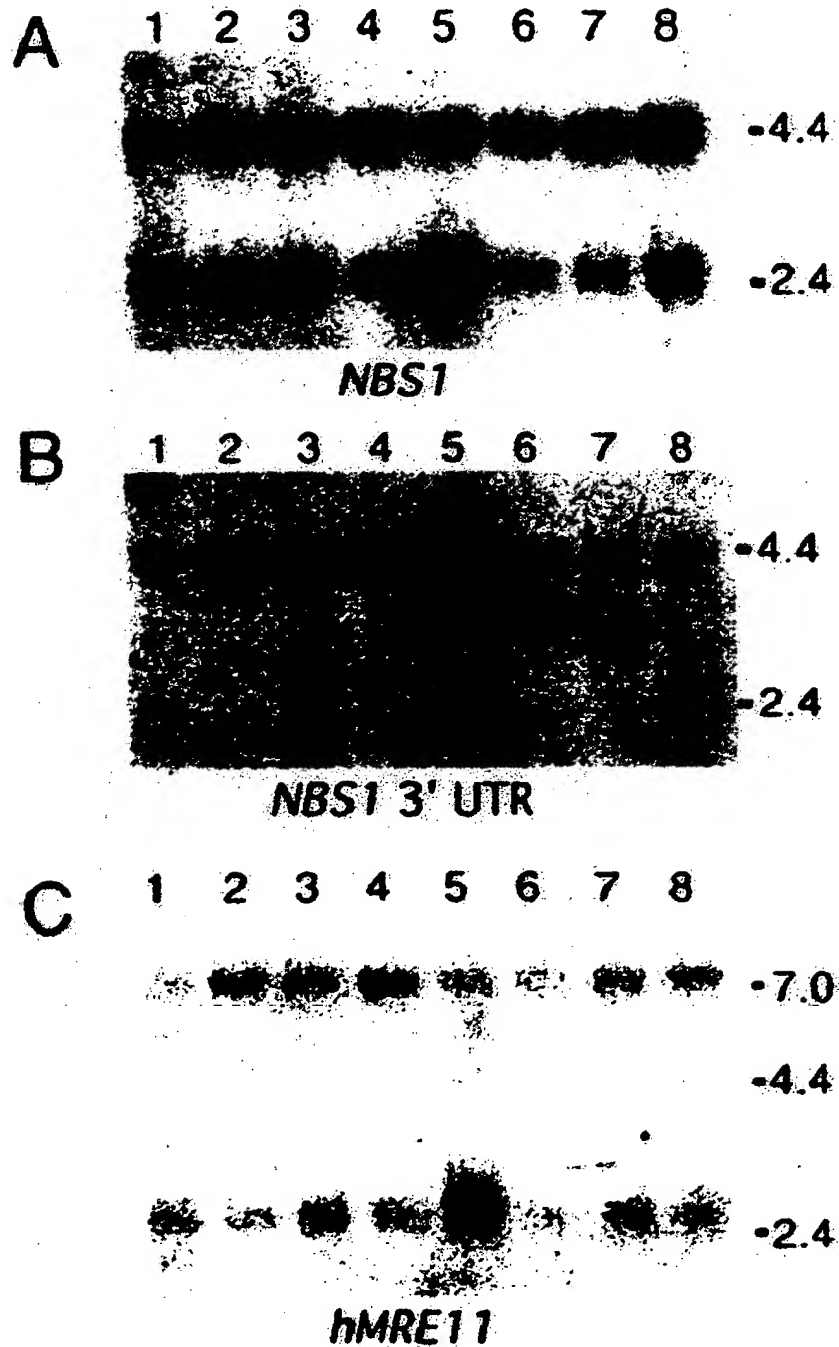


FIG. 7

7/13

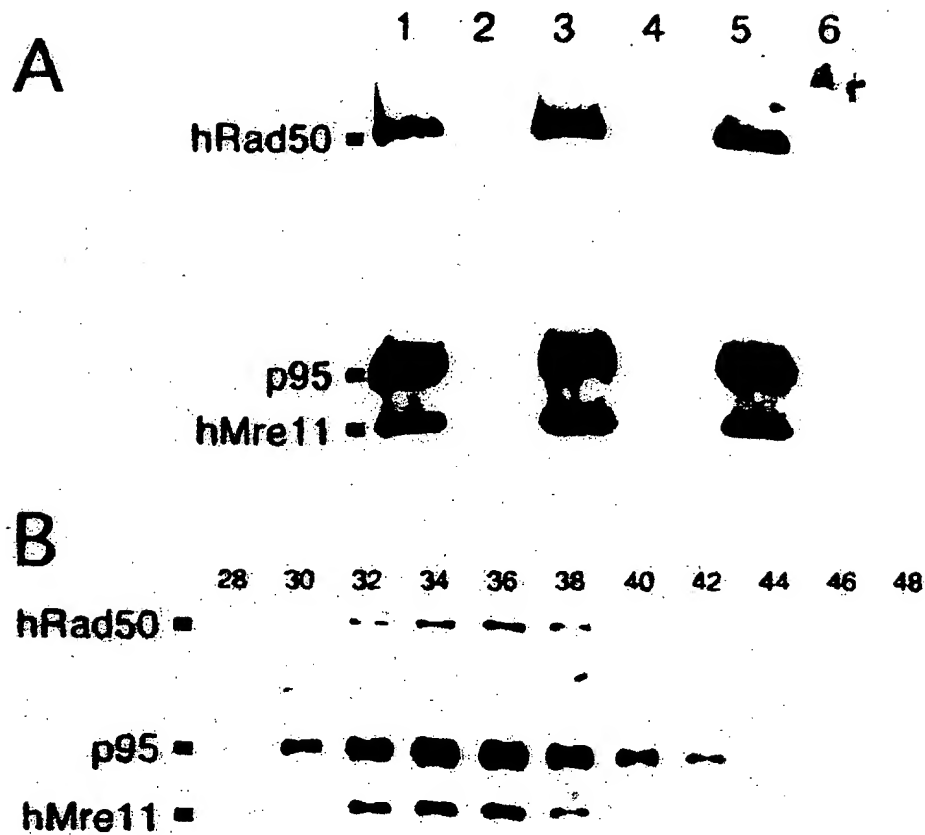


FIG. 8

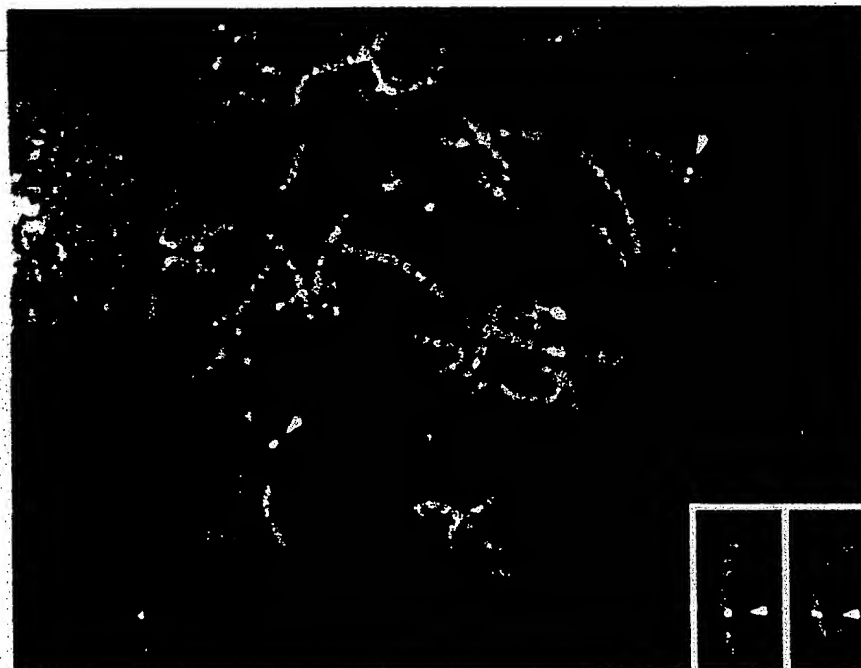


FIG. 9

8/13

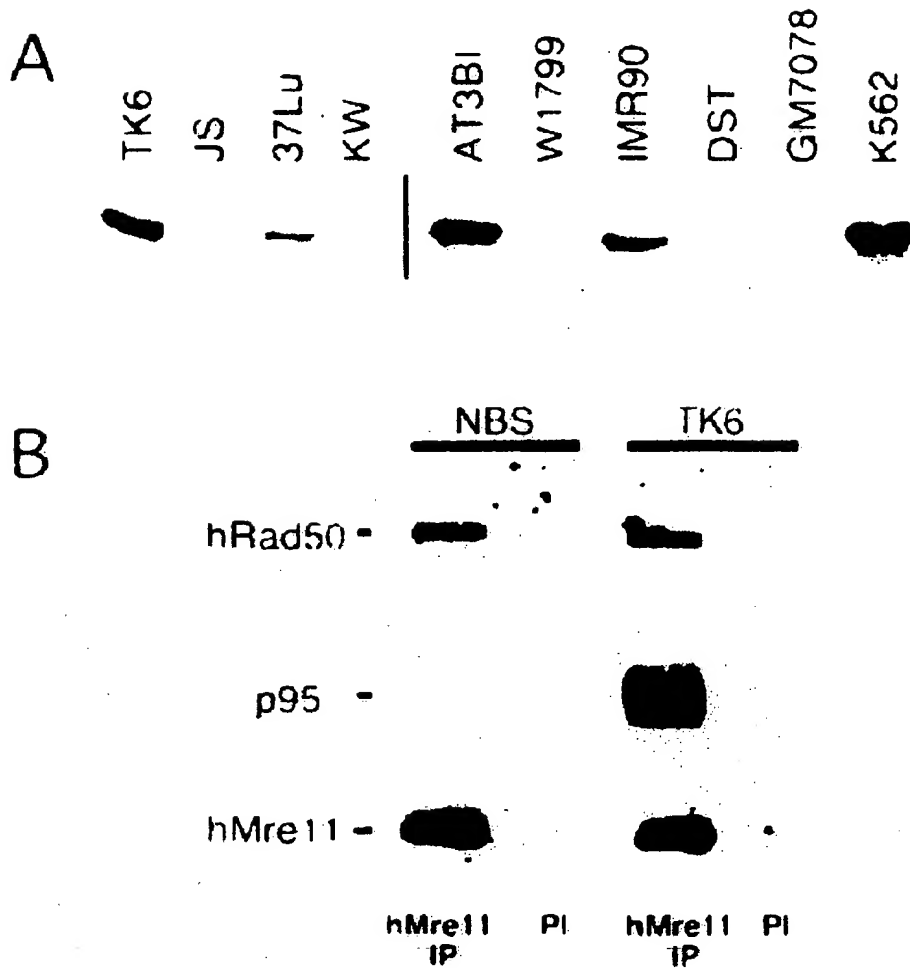


FIG. 10

9/13

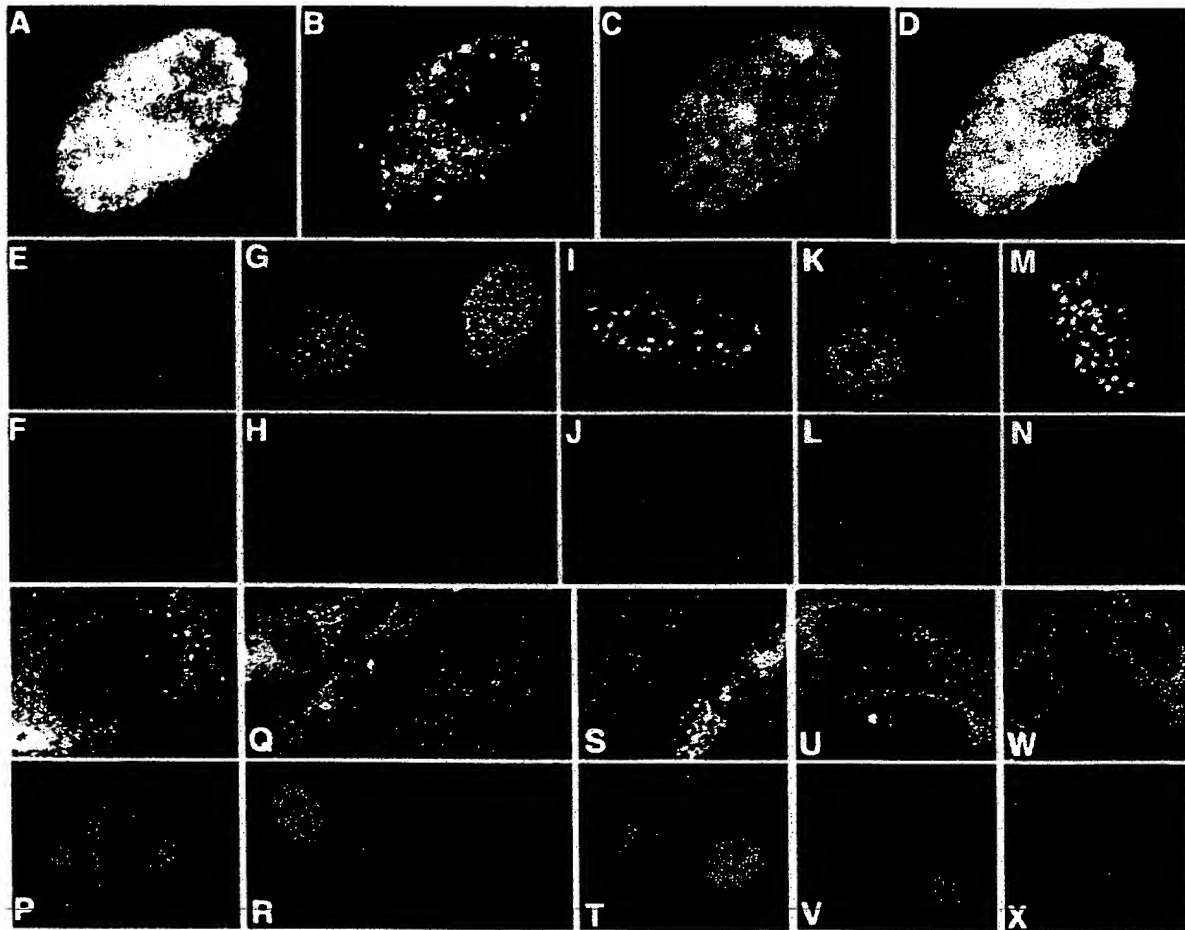


FIG. 11

10/13

<u>Amino Acid</u>	<u>Codon</u>
Phe	UUU, UUC
Ser	UCU, UCC, UCA, UCG, AGU, AGC
Tyr	UAU, UAC
Cys	UGU, UGC
Leu	UUA, UUG, CUU, CUC, CUA, CUG
Trp	UGG
Pro	CCU, CCC, CCA, CCG
His	CAU, CAC
Arg	CGU, CGC, CGA, CGG, AGA, AGG
Gln	CAA, CAG
Ile	AUU, AUC, AUA
Thr	ACU, ACC, ACA, ACG
Asn	AAU, AAC
Lys	AAA, AAG
Met	AUG
Val	GUU, GUC, GUA, GUG
Ala	GCU, GCC, GCA, GCG
Asp	GAU, GAC
Gly	GGU, GGC, GGA, GGG
Glu	GAA, GAG

FIG. 12

11/13

Original Residue	Exemplary Substitutions	Preferred Substitutions
Ala (A)	val; leu; ile	val
Arg (R)	lys; gln; asn	lys
Asn (N)	gln; his; lys; arg	gln
Asp (D)	glu	glu
Cys (C)	ser	ser
Gln (Q)	asn	asn
Glu (E)	asp	asp
Gly (G)	pro	pro
His (H)	asn; gln; lys; arg	arg
Ile (I)	leu; val; met; ala; phe norleucine	leu
Leu (L)	norleucine; ile; val; met; ala; phe	ile
Lys (K)	arg; gln; asn	arg
Met (M)	leu; phe; ile	leu
Phe (F)	leu; val; ile; ala	leu
Pro (P)	gly	gly
Ser (S)	thr	thr
Thr (T)	ser	ser
Trp (W)	tyr	tyr
Tyr (Y)	trp; phe; thr; ser	phe
Val (V)	ile; leu; met; phe; ala; norleucine	leu

FIG. 13

12/13

ttcggcacgaggcgcggttgcaagtcggccccagccctgaggagccggaccgatgtggaaactgctgcccgcgcgggc
cggcaggaggagaaccatacagacttttgactggcgttgagtacgttggttggaaggaaaaactgtgccattctaattgaa
aatgatcagtcgatcagccgaaatcatgctgtgttaactgctaacttttctgtaaccaacctgagtcacaacagatgaaa
ccctgtattgacattaaaagataattctaagtatggtaccttggtaaatgaggaaaaatgcagaatggcttttccgaa
ctttgaagtcgggggatggattacttttggagtggttggaagtaaatcagaatagagtatgagccttgggtgcatgc
tcttcttggtttagatgtctctgggaaaactgctttaaatacaagatatattgcaacttggaggatttactgtaacaattg
gacagaagaatgcactcaccttgcctgcatggatcagtgaaagttaccattaaaacaatatgtgcaactcatttgggacgtc
caattgtaaagccagaatattttactgaattcctgaaagcagttcagtcacaagaagcagcctccacaaattgaaagttt
taccacctcttgatgaaccatctattggaagtaaaaatgttgatctgtcaggacggcaggaaagaaaaacaaatcttcaa
agggaaaaacatttatattttgaaatgccaaacagcataaagaaattgagttccgcagttgtcttggagggtggggaagcta
gggtgataacagaagagaatgaagaagaacataatttcttttggctcgggaacgtgtgttggatgatacaggaataaca
aactcacagacctaattcctgactgtcagaagaatggattcagtcacaataatggatatgctccaaaggcaaggtcttag
acctattcctgaagcagaatggattggcgggtgatttcatgactacaaagaattactgtgatcctcagggccatccca
gtacaggattaaagacaaccaactccaggaccaagcctttcacaaaggcgtgtcagttgatgaaaaactaatgccaagcgc
ccagtgaaactacaacatacgtagctgacacagaatcagagcaagcagatacatgggatttgagtgaaggccaaaaga
aatcaaaagtctccaaatggaacaaaaattcagaatgctttcacaaagcgcacccactgtaaaggagtcctgcaaaacaa
gctctaataataatagtatggtatcaaaactttggctaagatgagaatcccaactatcagctttcaccaactaaatg
ccaagtataataaaaagtaagatagggcttctcagcagcagcagaccaactccatcagaaactactttcagcgtctac
caaaaaaagggaagggaatgaagaaatcaagaaatgtcttcatgcaaatcagcaagaatagaaacgtcttgttctctt
tagaacaacacaacctgctacacctcatttggtgaaaaataaggagcagcatctatctgagaatgagcctgtggacaca
aactcagacaataacttatttacagatcacagatttaaaatctattgtgaaaaattctgccagtaaatctcatgctgcaga
aaagctaagatcaataaaaaaagggaatggatgatgtggccatagaagatgaagtattggaacagttattcaaggaca
caaaaccagagttagaaattgatgtgaaagtcaaaaaacaggaggaagatgtcaatgttagaaaaaggccaaggatggat
atagaacaaatgacactttcagtgatgaagcagtaaccagaaagttagcaaaaatctcagaagaaatgaaattgggaagaa
acgtgaactcaagggaagactcactatgggtcagctaaagaaatatttcaaatgacaaactcaggatgatagtgagatgc
ttcaaaaaaagctgttattgactgaatttagatcactgggtgattaaaaactctacttccagaaatccgtctggcataaat
gatgattatgggtcaactaaaaaatttcaagaaattcaaaaaagggtcacatatcctggagcaggaaaaacttccacacatcat
tgaggatcagatctaataagctcatcatgctcgaagaatacagaactagaagagtggttaaggcaggaaatggaggtac
aaaatcaacatgcaaaagaagagtccttctgctgatgatctttttagatacaatccttatttaaaaaggagaagataac
aggattttaaagaagccatggaaaaacttccatagtaagcatctacttcaggccaacaagggttatatgaatatatagtg
tatagaagcgatttaagttacaatgttttatggcctaataatttataaataaaaatgcacaaaactttgattcttttga
taacaaattgttggtyctgttttcaggcttgcactaaagaaatatttcaaatgacaaactcaggatgatagtgagatgc
taatatagtcacagttcaaaattctaaatrtacgtaaggtaaggactaaagtcacccctccaccattgtcctagctact
tggttcccctcagaaaaaaattcatggatactcatttcttatgratcttccagggtttttgagtcctattcaaaatcc
tatttttaataatttccctacacaaatgatagcataacatatgcagtggtctacaccttgcttttttacttagtaaga
aaaaattataggaatatcaatataatgttttaatatatttttctttccattatgctgtagtcctacctaactctggtg
atccaaacaaaatggcttcagtggtgcagatgtcacctacatgttattctagtactagaaactgaagaccatgtggagac
ttcatcaaacatgggttttagttttcaccagaatggaaagacctgtaccccttttgggtggtcttactgagctgggtggg
gtcgtgttttgagcttatttagagtcctagtttctacttataaagtagaaatgggtgagattgtttcttttctacck
aaaggagatggtaagaaacaatgaatgtctttttcaaaacttattgacaagtgattttcaagtcgtgttcaaaaa
tattcatgtacctgtgatccagcaagaaggagttccagtcagagtcactacaactgattagttgttttagagaatgaga
aatggaacagtgaggaatggaggccatatttccatgacttccctgtaaacagaagcaacagaagggaagagggtggc
ctctacatcactctcaccttccaaatcttgggaagtgcacttacttgcagaaacaaattaacttacttccaagttctg
gctgcttgaggtggaactccagctgcaaggaggttagggaaatgaaggtctttttttaaagcttctcagccttccag
ggaacagaatgggtgagccaatctgcaatttctactacaggcattgagaccagttagattattgaaatattatagaga
gttatgaacacttaaattatgatagtggtatgacattggatagaacatgggatactttagaagtagaattgacagggcat
attagttgatgaaatggagtcatttgagtcctytaaagtcattgatacattaccaagtagaagtcgtggaacata
gtctccattttacagtttaaggaatataatggacagattaatatgttctgtctgcatgccacaatcccttctaaggaa
actgcctactatagcagtttttatatttgcatttatgaatataatgaatgaggagttctgggtacctcctgtctt
aaatattgggtgtgtccagtttttcccttttaacmctcccaattcgggtgtgtaggtggatgtttccatttgggt
tttaatttgtatatccctgatagctataattgggtcatagaatcttttatacattctagatgcaagtccttgyoggat
atacgtattgagatattacacctagtcgtggcttgactgttttctttatgtcttttgatgaatagaagttttaaatt
gacaaggtcaaaatttatttttttcttttgtttgatatttttctctccaaatttaaccccaagatttcagatatctg
tattatataaaactttatatttttatatttggatctacctgaattgatgtatgttgtgaattatggatcagggtct
tttttccccatacaagtatccagtcattgtaacactgtttattgaaagaattatccttccctcattaaattaccttgc
caattagtaaaaaatcaattaaccatrmarmmmrrrggatccactagttctagagcggccgccaccgcggtggagctcca
gct

FIG. 14

13/13

MWKLLPAAGPAGGEPYRLLTGVEYVVGKNCALLIENDQSI SRNHAVLTANFSVTNLSQTDEIPVLTLDNSKYGTFVNE
EKMONGFSRTLKSGDGITFGVFGSKFRIEYELVACSSCLDVSGKTALNQAILQLGGFTVNNWTEECTHLMVSVKVTIK
TICALICGRPIVKPEYFTEFLKAVQSKKQPPQIESFYPPLEPSIGSKNVDLSGRQERKQIFKGKTFIFLNAKQHKLLSS
AVVFGGGEARLITEENEEHNFFLAPGTCVVDGTGITSQTLIPDCQKKWISIMDMLQRQGLRPIPEAEIGLAVIFMTTK
NYCDPQGH PSTGLKTTTTPGPSLSQGVSVDEKLMPSAPVNTTTYVADTESEQADTWDL SERPKEIKVSKMEQKFRMLSQDA
PTVKESCKTSSNNNSMVSNTLAKMRI PNYQLSPTKLPSINKSKDRASQQQQTNSIRNYFPSTKKRERDEENQEMSSCKS
ARIETCSLLEQTQPATPSLWKNKEQHLS ENEPVD TNSDNNLFTD TDLKSIVKNSASKSHAAEKLRSNKKREMDDVAIED
EVLEQLFKDTKPELEIDVKVQKQ EEDVNVRKRPRMDIETNDTFSDEAVPESKISQENEIGKKRELKEDSLWSAKEISNN
DKLQDDSEMLPKKLLLTEFRSLVIKNSTSRNPSGINDDYQ LKNFKKFKKVTPGAGKLPHIIGGSDLIAHHARKNTELE
EWL RQEME VQNHAK EESLADDLFRYNPYLKRRR.

FIG. 15